SOFTWARE TOOL ARTICLE

Cytoscape tools for the web age: D3.js and Cytoscape.js exporters [version 1; peer review: 1 approved, 1 approved with reservations]

Keiichiro Ono, Barry Demchak, Trey Ideker
Department of Medicine, University of California, San Diego, La Jolla, CA 92093-0688, USA

Abstract
In this paper we present new data export modules for Cytoscape 3 that can generate network files for Cytoscape.js and D3.js. Cytoscape.js exporter is implemented as a core feature of Cytoscape 3, and D3.js exporter is available as a Cytoscape 3 app. These modules enable users to seamlessly export network and table data sets generated in Cytoscape to popular JavaScript library readable formats. In addition, we implemented template web applications for browser-based interactive network visualization that can be used as basis for complex data visualization applications for bioinformatics research. Example web applications created with these tools demonstrate how Cytoscape works in modern data visualization workflows built with traditional desktop tools and emerging web-based technologies. This interactivity enables researchers more flexibility than with static images, thereby greatly improving the quality of insights researchers can gain from them.

This article is included in the Cytoscape Apps gateway.

This article is included in the International Society for Computational Biology Community Journal gateway.
Corresponding author: Keiichiro Ono (kono@ucsd.edu)

Competing interests: No competing interests were disclosed.

Grant information: This work was funded by the National Resource for Network Biology (NRNB) under award numbers P41RR031228 and GM103504 assigned to Trey Ideker.

Copyright: © 2014 Ono K et al. This is an open access article distributed under the terms of the Creative Commons Attribution Licence, which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited. Data associated with the article are available under the terms of the Creative Commons Zero "No rights reserved" data waiver (CC0 1.0 Public domain dedication).

How to cite this article: Ono K, Demchak B and Ideker T. Cytoscape tools for the web age: D3.js and Cytoscape.js exporters [version 1; peer review: 1 approved, 1 approved with reservations] F1000Research 2014, 3:143 (https://doi.org/10.12688/f1000research.4510.1)

First published: 01 Jul 2014, 3:143 (https://doi.org/10.12688/f1000research.4510.1)
Introduction
Cytoscape was born as a GUI-based, Java desktop application in 2003\textsuperscript{1,2}. Today, it is a de-facto standard application for biological network analysis and visualization. Around 2005, Java was one of the dominant programming languages for data visualization applications, and Java-based feature-rich toolkits were developed\textsuperscript{3}. However, since they were designed before the re-discovery of Ajax\textsuperscript{4}, developers could not predict the success of JavaScript and related web technologies today. Cytoscape is still an important platform for biological network data integration and analysis, but for data visualization and sharing, we need a new method to take advantage of modern web technologies. Utilizing HTML5 and other emerging web technologies, Cytoscape Consortium developed a JavaScript library for network visualization called cytoscape.js (http://cytoscape.github.io/cytoscape.js/), the successor of Cytoscape Web\textsuperscript{5}, to meet the demand from the Cytoscape user community. Although Cytoscape and Cytoscape.js share some of the core concepts, such as Visual Styles or automatic layouts, they are completely independent software packages and there has been no simple way to use Cytoscape data sets in Cytoscape.js.

Public biological data repositories are still growing rapidly and the demand for visualizing those complex biological data sets is high. Traditionally, analysis and visualization of biological data is done by desktop applications, and in most cases, visualizations created by popular libraries (matplotlib\textsuperscript{6}, ggplot\textsuperscript{2}) are static images. It is hard to perform exploratory data analysis only with static images, especially when visualizing large data sets, because some of the details are lost due to the limited size of printed papers or computer screens. Instead of developing custom visualization toolkits for specific data sets, the scientific data visualization community is heading towards web-based technologies to utilize actively developed visualization toolkits such as mpld3 (http://mpld3.github.io/) or Bokeh (http://bokeh.pydata.org/). D3.js\textsuperscript{7} is one of the most popular toolkits for creating custom interactive visualizations. If biologists can use the existing powerful desktop application and these emerging web technologies for data visualizations and sharing, it opens up a new way to understand large and complex biological data sets.

To bridge the gap between the desktop version of Cytoscape and other web-based data visualization toolkits, we developed Cytoscape modules to generate web-friendly data formats. The goal is to enable shared visualization of Cytoscape networks via a web platform (e.g., browser), and our strategy is to enable conversion from Cytoscape data objects to a format friendly to web apps, and to provide a template code for creating an interactive web application. In this paper, we present the implementation of Cytoscape data exporters and template web applications and demonstrate how users can publish their data sets as interactive data visualizations with our new tools.

Implementation
The exporter modules were developed for Cytoscape 3. The Cytoscape.js exporter is part of Cytoscape 3 core distribution and is available as a standard feature. The D3.js exporter is an app, and this means developers can write any type of additional JSON exporters as necessary. Exporter modules generate JavaScript Object Notation (JSON) files that are readable by Cytoscape.js and D3.js. To visualize these JSON files as interactive network diagrams, users have to write some JavaScript and HTML5 code to read, map, and render the network and table data in the files. The general structure of code for basic network visualization is common to most use cases. To simplify this visualization task, we implemented template HTML5 projects to render the exported JSON files.

Cytoscape.js exporter
Cytoscape.js is a JavaScript library for interactive network visualization developed by Cytoscape Consortium. Although Cytoscape and Cytoscape.js share core concepts, they have completely independent code bases written in Java and JavaScript. This means there is no binary-level compatibility between these two software packages. The purpose of this exporter is to provide data-level compatibility between these software packages. There are two core functions in this exporter module: network/table converter and Visual Style to CSS converter.

Network/Table to JSON converter
Conversion from Cytoscape networks and tables to Cytoscape.js JSON is done by a serialization module implemented with Jackson, a Java-based JSON parser library (https://github.com/FasterXML/jackson). The converter takes a Cytoscape network object and associated node, edge, and network tables as inputs and converts them into a single JavaScript object represented as JSON. Most of the basic data types are converted into JSON except the following: nested networks, custom graphics, and node shapes and edge line types, which are only available in Cytoscape.

Visual style to CSS
In contrast, converting Cytoscape Visual Style is a nontrivial process. A Visual Style in Cytoscape is a collection of visual mapping functions, which is a mapping from data to visual variables\textsuperscript{8}, and default values. Conversion from default visual property values to JavaScript objects is a simple one-to-one mapping with several non-compatible value filterings. In Cytoscape, there are three types of visual mapping functions. They are Passthrough, Continuous, and Discrete. Cytoscape.js has the concept of a visual mapping function in its design and it follows standards of CSS and selectors, which is significantly different from the design of Cytoscape visual mapping functions. The converter translates Cytoscape visual mapping functions into combinations of Cytoscape.js selectors and mappers (Figure 1). This translation absorbs differences in design between the two applications and reproduces Cytoscape Visual Styles as JavaScript objects used for styling in Cytoscape.js.

D3.js exporter
In general, D3.js does not have any specific data format for visualization. Generic CSV/TSV tables can be used for all types of visualizations, and its core provides data loaders for those files. An exception is the graph data format for force-layout, which is the basic preset for visualizing graph data in D3.js (https://github.com/mbostock/d3/wiki/Force-Layout). It uses an ordinal (i.e., zero-based) index of nodes as the unique identifier, and edges are represented as a pair of those indices. The D3.js exporter converts Cytoscape network topology into this force-layout format, and transforms all associated data tables into properties of nodes and edges in the JSON.
A tree data structure is a special kind of graph, and D3.js has various types of preset visualizations for it, such as radial layout, circle packing or Treemap. Cytoscape can visualize trees as node-link diagrams, and if we can export tree data models stored as Cytoscape graph objects into a D3.js compatible format, users can create multiple views for same data sets using different visualization techniques which could provide new insights for them. To utilize these presets, the exporter generates tree-style JSON for D3.js. The root node of the tree should be specified manually by the user, then the exporter automatically generates tree JSON with all associated tables.

Template projects for visualizing JSON exports
Once exporters generate JSON files, users need actual web applications to visualize the data. Both Cytoscape.js and D3.js are designed for developers, not for end-users, and developers are expected to write their own custom visualization code to see the data. Although they are optimized for custom web-based visualizations, basic components of visualization code, including data loading, mapping, and rendering, are common to most applications. To minimize duplicate efforts to visualize the results from JSON exporters, we developed template web application projects to visualize JSON files generated by Cytoscape. These templates can create basic visualizations of the JSON files out of the box. To develop these templates, we used standard tools for modern JavaScript development: Node.js (http://nodejs.org/) as runtime for all development tools, Yeoman (http://yeoman.io/) for code scaffolding, and Grunt (http://gruntjs.com) as task runner.

Results
A typical data visualization workflow with our new tools consists of the following four steps. First, users load networks, annotations, and experimental data sets into Cytoscape. Second, utilizing core Cytoscape visualization features, users create custom Visual Styles and layouts. Third, all data sets are exported as JSON files, and finally they can create custom web-based, interactive visualizations from the template projects (Figure 2). The original network data visualized in Figure 3 was imported with a Cytoscape app called KEG-GScape (http://apps.cytoscape.org/apps/kegggscape). The advantage of this workflow is that users can use the large collection of existing Cytoscape apps for data integration and analysis, and the result can be exported as interactive web-based visualizations with the new exporters. Figure 3 shows the TCA Cycle pathway which was generated from a KEGG XML (KGML) file, and its Visual Style was automatically generated from the graphics data in the file. Cytoscape.js exporter can generate web-compatible style and network files directly from the Cytoscape view. Our template code for Cytoscape.js is a simple viewer, and it can be used as a basis for complex data visualization application.

The exporters create JSON files with both network topology and data tables, and users can create complex data visualizations which cannot be achieved with Cytoscape alone. Figure 4 and Figure 5 shows simple network visualizations created with D3.js force-directed layout (Figure 4) and tree layout (Figure 5). These figures are created with a minimal set of features available in D3.js and they can be used as a “boilerplate” code for custom visualizations. The desktop version of Cytoscape is optimized for rendering node-link, or ball-stick network diagrams, which is only one way to visualize graph data. Cytoscape 3 supports multiple-rendering engines and if developers can implement new rendering engines for new visualizations, such as Treemap or Chord Diagram, they can add new visualizations on Cytoscape. This is not a trivial task and as an alternative, our D3.js templates can be used to make prototypes for new visualizations.
Figure 2. Basic workflow for publishing Cytoscape-generated networks as interactive web visualizations. The combination of Cytoscape 3 and new JSON exporters can be used as data integration tool for web-based visualizations. Our sample Grunt project generates a simple template code for visualizing D3.js and Cytoscape.js JSON files.

Figure 3. Sample view of exported JSON file visualized by the template code using Cytoscape.js. Original network data was imported to Cytoscape 3 using the KEGGscape app. The template code contains minimal set of features like simple table browser and network viewer. The table browser (bottom) is implemented with AngularJS (https://angularjs.org) and Bootstrap (http://getbootstrap.com).
Figure 4. Visualization of a sample network (galFiltered.sif) by D3.js force-layout.

Figure 5. Tree version of Gene Ontology visualized by D3.js tree layout.
Conclusions
In this paper, we presented a new workflow to visualize biological data sets using Cytoscape and modern web-based data visualization libraries. The example visualizations show how users can leverage easy-to-use Cytoscape core features as a part of web-based interactive data visualization publishing workflow.

Future plan
At this point, new features discussed in this paper are designed for developers who can write JavaScript and HTML5 code. End users are also a part of our target audience, and so we will implement the “Export as HTML5 Session” feature as a core Cytoscape feature, which creates a compressed archive file that includes all of the networks, tables, and Visual Styles as JSON along with all JavaScript files to visualize the data as a single-page application.

Software availability
Software available from: http://apps.cytoscape.org/apps/d3jsexporter
Latest source code: https://github.com/keiono/cytoscape-d3
Source code as at the time of publication: https://github.com/F1000Research/cytoscape-d3/releases/tag/V1.0
Archived source code as at the time of publication: http://dx.doi.org/10.5281/zenodo.1054710

License MIT License
Template code generator website https://github.com/cytoscape/cyjs-sample
High-resolution images and interactive examples are available at our web sites above.

Author contributions
KO designed and implemented all software packages and wrote this article. BD worked as a project manager and reviewed this article. TI provided funding and supervision.

Competing interests
No competing interests were disclosed.

Grant information
This work was funded by the National Resource for Network Biology (NRNB) under award numbers P41 RR031228 and GM103504 assigned to Trey Ideker.

Acknowledgements
We would like to thank Max Franz and Dr. Gary Bader for their work on the Cytoscape.js JavaScript library.

References

Open Peer Review

Current Referee Status: ? ✔️

Version 1

Referee Report 11 September 2014
https://doi.org/10.5256/f1000research.4824.r6061

Jose Villaveces
Research Group Computational Biology, Max Planck Institute of Biochemistry, Munich, Germany

The article describes Cytoscape.js and D3.js exporters and a code template generator. Additionally, it provides all the essential information about Cytoscape.js exporter and D3.js exporters and points interested users to the source code for its implementation.

Even though there are a few points that require clarification, the tools are a nice addition to Cytoscape and can be seen as a first step in uniting present day web technologies and the more conventional desktop based visualisation provided by Cytoscape.

Improvements
1. There is no documentation for the template generator, a README.md file with a few examples will be nice.

2. The D3 example listed in the d3jsexporter page is very small (300 x 150) and the 'View Source' button is not working.

Competing Interests: No competing interests were disclosed.

I have read this submission. I believe that I have an appropriate level of expertise to confirm that it is of an acceptable scientific standard.

Referee Report 01 August 2014
https://doi.org/10.5256/f1000research.4824.r5313

Alexander Lex
Visual Computing Group, Harvard University, Cambridge, MA, USA

The article describes an important new development for the Cytoscape project: bridging the gap between complex yet powerful desktop applications to easy-to-use, installation free web application focused on the analysis and presentation. I consider this workflow as an important bridge between the two worlds. The article is technically sound and mostly well written. Some structural comments and concrete suggestions for improving the manuscript are listed below.
What I am currently missing in the manuscript is a description of a workflow for cleaning up the exported visualizations. Especially in the KEGG example, the result is rather messy and clearly requires some additional authoring. How would one go about this process? How long would it take? I would suggest adding another figure that shows a post-cleanup version and a paragraph commenting on the process.

Also, it would be great to see a comparison of the original layout in Cytoscape in addition to the web-based layouts, ideally for all the results figures, so that readers can clearly judge the quality of the approach. This could also put my previous comment into context, as maybe the KEGG layout isn't much better in Cytoscape standalone, since the KEGG annotations typically don't contain sufficient information about the layout.

**Structural Suggestions:**

- For this argument:

  "However, since they were designed before the re-discovery of Ajax, developers could not predict the success of JavaScript and related web technologies today."

  I would suggest to add that it was simply not feasible at the time since the technology and bandwidth was not available.

- Regarding this sentence:

  "It is hard to perform exploratory data analysis only with static images, especially when visualizing large data sets, because some of the details are lost due to the limited size of printed papers or computer screens."

  I would suggest to also argue that this static approach is especially problematic for networks, as the size of all but trivial networks requires dynamic features to be useful.

- Here:

  "If biologists can use the existing powerful desktop application and these emerging web technologies for data visualizations and sharing, it opens up a new way to understand large and complex biological data sets."

  I think the main point is that it has the potential to combine the best of two worlds: powerful authoring tools and easily shareable interactive results, but I'm not sure whether this is "a new way to understand[ing]". Rather it is a new way of engaging users, through a much simplified process.

- "via a web platform (e.g., browser)" -> what would be other web platforms? Or do you mean "i.e., in the browser"

- The hierarchy in the Cytoscape.js exporter section isn't clear. The following two captions seem to be of the same level as the root section. I suggest removing the captions completely and simply make the converters bold in the text. Also, these sections sometimes lack clarity. For example, it is not clear what a Cytoscape Table and Network is. I assume that is clear when one knows the
software well, but it would be good to clarify this here for readers who are not that familiar with Cytoscape.

- What are "non-compatible value filterings"?
- "The root node of the tree should be specified manually by the user" -> should or must?
- Caption Figure 4: "Visualization of a sample network (galFiltered.sif) by D3.js force-layout." -> What is the data shown here? Should a reader know that from the file name? The file name seems irrelevant here. I would rather add a brief description of the data.

Some grammar and style suggestions:
- "to popular JavaScript library readable formats" -> to "formats readable by popular JavaScript libraries."
- "and Java-based feature-rich toolkits were developed" -> "and many Java-based feature-rich toolkits were developed".
- "technologies, Cytoscape Consortium" -> "technologies, the Cytoscape Consortium".
- "libraries (matplotlib, ggplot2)" -> "libraries (e.g., matplotlib, ggplot2)".
- "The goal is to enable shared visualization of Cytoscape networks via a web platform (e.g., browser), and our strategy is to enable conversion from Cytoscape data objects to a format friendly to web apps, and to provide a template code for creating an interactive web application." -> split into multiple sentences.
- "is part of Cytoscape 3 core distribution" -> "is part of the Cytoscape 3 core distribution".
- "The D3.js exporter is an app, and this means" -> "The D3.js exporter is an app which means"
- "The D3.js exporter is an app, and this means" -> I suggest clarifying "app". Is this a cytoscape plugin?
- "we implemented template HTML5 projects to render the exported JSON files." -> "we provide template HTML5 projects to render the exported JSON files.".
- "developed by Cytoscape Consortium" -> "developed by the Cytoscape Consortium".
- "There are two core functions in this exporter module: network/table converter and Visual Style to CSS converter." -> I would suggest rephrasing this to explain on a high level what these modules do.
- "except the following: nested networks," -> "except for nested networks,".
- "converting Cytoscape Visual Style" -> is this a noun? or do you mean "Cytoscape's visual styles".
In Figure 1, the code in the "Continuous Mapping as collection of Mappers" is unreadable as it is too small.

"Once exporters generate JSON files, users need actual web applications to visualize the data." -> "To visualize the exported JSON files, actual web applications are needed to visualize the data.".

"visualization code to see the data." -> "visualization code."

"Third, all data sets are exported as JSON files, and finally they can create custom web-based, interactive visualizations from the template projects (Figure 2)." -> This sentence mixes active and passive. I suggest re-writing it in the active voice since the previous sentence is active as well.

"The combination of Cytoscape 3 and new JSON exporters can be used as data integration tool for web-based visualizations" -> "The combination of Cytoscape 3 and the new JSON exporters can be used as a data integration tool for web-based visualizations."

"created with D3.js force-directed layout" -> "created with the D3.js force-directed layout".

"such as Treemap or Chord Diagram" -> "such as Treemaps or Chord Diagrams".

"License MIT License" -> "License: MIT License".


**Competing Interests:** No competing interests were disclosed.

I have read this submission. I believe that I have an appropriate level of expertise to confirm that it is of an acceptable scientific standard, however I have significant reservations, as outlined above.

**Author Response 06 Aug 2014**

Keiichiro Ono, University of California, San Diego, USA

I completely agree with your point about the missing details for web based KEGG visualization part. I will add more details how to reproduce the visualization with this tool. It will be available as a GitHub Wiki page. The following details will be added to the new version:

- Side-by-side comparison of original Cytoscape visualization and exported version
- Original Cytoscape Session file and exported JSON files as supplemental data
- Link to the instruction wiki page how to reproduce the KEGG visualization

In addition to these changes, I will fix the structural/grammatical issues in the revised version.

Thank you for your helpful suggestions.

**Competing Interests:** No competing interests were disclosed.

**Author Response 25 Nov 2014**

Keiichiro Ono, University of California, San Diego, USA
Dear Dr. Lex,

This version 2 includes all updates for your comments for the original version. Also, recently we released new version of Cytoscape (3.2.0), and it should be compatible with this protocol. If you have problems to test this workflow in the latest environment, please let me know.

Best,
Keiichiro Ono

**Competing Interests:** No competing interests were disclosed.